**TDMS No.** 96011 - 05

Test Type: CHRONIC

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

**Species/Strain:** RATS/F 344 **Pathologist:** PIERCE, J. - RENNE, R.

J. - KEININE, K. Lab. I

F 1 R2

**C Number:** C96011

Route: RESPIRATORY EXPOSURE WHOLE BODY

**Lock Date:** 01/28/2004

Cage Range: ALL

Date Range: ALL

**Reasons For Removal:** ALL

Removal Date Range: ALL

Treatment Groups: Include ALL

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS MALE	CONTROL	250 PPM	500 PPM	1000 PPM
Disposition Summary				
Animals Initially in Study	50	50	50	50
Early Deaths				
Moribund Sacrifice	20	24	21	24
Natural Death	4	3	2	2
Survivors				
Terminal Sacrifice	26	23	27	24
Animals Examined Microscopically	50	50	50	50
ALIMENTARY SYSTEM				
Esophagus	(50)	(50)	(50)	(50)
Foreign Body	()	1 [4.0]	(,	()
Intestine Large, Cecum	(49)	(49)	(49)	(49)
Intestine Large, Colon	(49)	(50)	(50)	(50)
Diverticulum	1 [2.0]	, ,		, ,
Intestine Large, Rectum	(49)	(50)	(50)	(50)
Intestine Small, Duodenum	(49)	(49)	(49)	(49)
Intestine Small, Ileum	(49)	(47)	(48)	(49)
Intestine Small, Jejunum	(48)	(47)	(48)	(49)
Inflammation, Chronic Active	1 [4.0]			
Thrombosis	1 [4.0]			
Ulcer	1 [4.0]	(=o)	(=o)	(50)
Liver	(50)	(50)	(50)	(50)
Angiectasis	7	1 [2.0]	0	C
Basophilic Focus Clear Cell Focus	7 12	4 6	8 11	6 11
Degeneration, Cystic		_	4 [2.5]	6 [3.0]
Eosinophilic Focus	3 [1.0]	1 [3.0] 1	4 [2.5] 3	ช [อ.บ]
Eosinophilic Focus, Multiple		ļ	3 1	
Fatty Change, Diffuse		1 [3.0]	ı	
Hepatodiaphragmatic Nodule	3 [4.0]	8 [4.0]	2 [4.0]	3 [4.0]
Inflammation, Suppurative	0[0]	1 [4.0]	- [·]	0[0]
Mixed Cell Focus	1	. [ ]	3	1
Necrosis	3 [4.0]	3 [2.7]	3 [3.0]	3 [2.3]
Vacuolization Cytoplasmic	6 [2.7]	2 [3.0]	3 [2.3]	2 [2.0]
Bile Duct, Hyperplasia	7 [2.4]	4 [2.3]	12 [3.2]	7 [2.7]

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

**CUMENE** 

**CAS Number: 98-82-8** 

Pathologist: PIERCE, J. - RENNE, R.

Date Report Regsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS MALE	CONTROL	250 PPM	500 PPM	1000 PPM	
Hepatocyte, Regeneration	1 [4.0]				
Oval Cell, Hyperplasia				1 [4.0]	
Periportal, Inflammation, Chronic			2 [3.5]		
Mesentery	(7)	(13)	(10)	(6)	
Hemorrhage		1 [2.0]		1 [3.0]	
Necrosis	6 [2.5]	10 [2.9]	10 [3.0]	3 [3.3]	
Fat, Hemorrhage	1 [3.0]			1 [3.0]	
Fat, Necrosis		1 [3.0]			
Pancreas	(50)	(50)	(50)	(50)	
Hyperplasia				1 [3.0]	
Vacuolization Cytoplasmic		1 [2.0]			
Acinus, Atrophy	2 [1.0]	2 [3.0]	2 [3.0]	2 [2.5]	
Acinus, Hyperplasia			1 [2.0]	3 [1.7]	
Stomach, Forestomach	(50)	(50)	(50)	(50)	
Hemorrhage	1 [2.0]				
Inflammation, Suppurative			2 [2.0]		
Necrosis	1 [2.0]				
Ulcer	3 [3.3]	7 [3.9]	1 [4.0]	8 [3.9]	
Epithelium, Hyperplasia	6 [2.7]	11 [3.0]	7 [3.4]	10 [3.4]	
Stomach, Glandular	(50)	(50)	(50)	(50)	
Erosion	2 [2.0]	1 [3.0]	1 [3.0]	2 [2.0]	
Hyperplasia, Lymphoid			1 [3.0]		
Mineralization	1 [4.0]				
Ulcer	3 [3.3]		1 [4.0]	1 [4.0]	
Tongue	(1)	(3)	(1)	(3)	
Epithelium, Hyperplasia	1 [3.0]	2 [3.5]	1 [3.0]	3 [2.0]	
Tooth	(0)	(1)	(0)	(0)	
Inflammation		1 [4.0]			
ARDIOVASCULAR SYSTEM					
Blood Vessel	(1)	(1)	(1)	(0)	
Mineralization	1 [4.0]				
Heart	(50)	(50)	(50)	(50)	
Cardiomyopathy	9 [1.8]	7 [2.6]	12 [2.0]	11 [1.7]	
Atrium, Thrombosis	3 [3.7]	6 [3.0]	4 [2.8]	4 [3.5]	
Valve, Cardiomyopathy	1 [3.0]				
Valve, Hemorrhage			1 [2.0]		
Valve, Thrombosis	1 [4.0]				

a - Number of animals examined microscopically at site and number of animals with lesion b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Regsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS MALE	CONTROL	250 PPM	500 PPM	1000 PPM	
Ventricle, Hypertrophy				1 [4.0]	
ENDOCRINE SYSTEM					
Adrenal Cortex Hyperplasia Hypertrophy Vacuolization Cytoplasmic Adrenal Medulla	(50) 17 [2.2] 8 [2.0] (50)	(50) 16 [2.7] 5 [2.2] (50)	(50) 10 [2.4] 1 [3.0] 7 [2.3] (50)	(50) 10 [2.7] 4 [2.5] (50)	
Hyperplasia Bilateral, Hyperplasia Islets, Pancreatic	11 [2.5] (50)	13 [2.5] (50)	(50) 18 [2.8] (50)	(50) 12 [2.5] 1 [2.0] (50)	
Hyperplasia Parathyroid Gland Hyperplasia Pituitary Gland	1 [3.0] (45) 1 [4.0] (50)	1 [3.0] (49) 1 [4.0] (50)	(49) (49)	(48) (50)	
Cyst Hemorrhage Hyperplasia Pars Intermedia, Hyperplasia	1 [4.0] 6 [3.5] 1 [2.0]	1 [3.0] 1 [4.0] 7 [2.7]	2 [3.5] 1 [4.0] 13 [3.2]	3 [4.0] 6 [3.0]	
Thyroid Gland C-cell, Hyperplasia Follicular Cell, Cyst Follicular Cell, Hyperplasia	(50) <sup>*</sup> 7 [1.9]	(50) 11 [1.8] 1 [2.0]	(50) 6 [1.8] 1 [2.0]	(50) 8 [2.0]	
GENERAL BODY SYSTEM					
Peritoneum	(2)	(2)	(4)	(2)	
GENITAL SYSTEM					
Epididymis Preputial Gland Cyst Hyperplasia	(50) (50) 1 [4.0]	(50) (50)	(50) (50) 2 [2.5]	(50) (50) 1 [4.0]	
Inflammation, Suppurative Prostate Hyperplasia	(50)	(50) 1 [2.0]	1 [4.0] (50) 1 [2.0]	(50) 2 [1.5]	

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS MALE	CONTROL	250 PPM	500 PPM	1000 PPM	
Inflammation, Suppurative	42 [1.6]	32 [1.7]	27 [2.0]	28 [1.7]	
Seminal Vesicle	(50)	(50)	(50)	(50)	
Dilatation	(00)	1 [4.0]	(00)	(88)	
Hyperplasia		1 [3.0]			
Testes	(50)	(50)	(50)	(50)	
Necrosis	()	()	1 [4.0]	()	
Artery, Inflammation, Chronic Active		1 [3.0]	2 [3.0]		
Bilateral, Interstitial Cell, Hyperplasia			[]	1 [2.0]	
Germinal Epithelium, Atrophy	6 [3.5]	5 [3.8]	3 [2.7]	6 [3.2]	
Interstitial Cell, Hyperplasia	12 [1.9]	18 [2.1]	19 [2.2]	9 [3.0]	
EMATOPOIETIC SYSTEM					
Bone Marrow	(50)	(50)	(50)	(50)	
Lymph Node	(7)	(10)	(8)	(14)	
Deep Cervical, Ectasia		1 [4.0]		1 [4.0]	
Deep Cervical, Hemorrhage				1 [3.0]	
Deep Cervical, Hyperplasia, Lymphoid			1 [2.0]	2 [3.5]	
Pancreatic, Ectasia			1 [2.0]	1 [3.0]	
Pancreatic, Hyperplasia, Lymphoid			1 [1.0]		
Pancreatic, Inflammation, Granulomatous			1 [2.0]		
Lymph Node, Bronchial	(7)	(12)	(11)	(8)	
Ectasia	1 [4.0]	1 [4.0]			
Hyperplasia, Lymphoid	1 [2.0]	1 [2.0]	1 [4.0]		
Lymph Node, Mandibular	(1)	(3)	(0)	(1)	
Ectasia	1 [4.0]	2 [4.0]			
Metaplasia, Osseous	1 [3.0]				
Lymph Node, Mediastinal	(34)	(32)	(34)	(34)	
Angiectasis				1 [4.0]	
Hemorrhage		1 [2.0]			
Hyperplasia, Lymphoid	1 [4.0]		2 [3.5]		
Inflammation, Suppurative	1 [4.0]	,	/·	(==)	
Lymph Node, Mesenteric	(50)	(50)	(50)	(50)	
Hemorrhage	(m-1	2 [3.5]	/·	1 [2.0]	
Spleen	(50)	(50)	(50)	(50)	
Accessory Spleen	0.50.71	0.10.01	7 (0.0)	1	
Fibrosis	2 [3.5]	2 [3.0]	7 [3.6]	2 [3.5]	
Hematopoietic Cell Proliferation	4.54.03	4.54.03	0.14.03	1 [3.0]	
Hemorrhage	4 [4.0]	4 [4.0]	2 [4.0]	4 [4.0]	

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Regsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS MALE	CONTROL	250 PPM	500 PPM	1000 PPM	
Necrosis Thymus	1 [3.0] (49)	3 [3.7] (49)	1 [4.0] (50)	(50)	
INTEGUMENTARY SYSTEM					
Mammary Gland Galactocele Epithelium, Hyperplasia Skin Cyst Epithelial Inclusion Hyperkeratosis Inflammation, Chronic Ulcer	(50) 2 [4.0] 1 [4.0] (50) 1 [4.0] 1 [3.0]	(50) (50) 1 [4.0]	(50) 2 [4.0] 1 [4.0] (50) 1 [4.0]	(50) 1 [2.0] 1 [2.0] (50) 2 [4.0] 1 [4.0] 1 [4.0]	
MUSCULOSKELETAL SYSTEM					
Bone Hyperostosis Skeletal Muscle	(50) (1)	(50) (2)	(50) 1 [3.0] (2)	(50) (1)	
NERVOUS SYSTEM					
Brain Compression Hemorrhage Necrosis Thrombosis	(50) 7 [3.6] 5 [3.6]	(50) 16 [3.1] 7 [2.3] 1 [2.0] 1 [2.0]	(50) 6 [2.7] 3 [2.0]	(50) 6 [3.2] 1 [2.0]	
RESPIRATORY SYSTEM					
Larynx Foreign Body Inflammation, Suppurative Inflammation, Chronic Epiglottis, Hyperplasia	(50) 6 [4.0] 4 [1.8] 1 [3.0] 1 [1.0]	(50) 3 [3.0] 7 [1.6] 1 [2.0]	(50) 1 [1.0] 1 [3.0]	(50) 3 [4.0] 3 [1.3]	
Epiglottis, Hyperplasia Epiglottis, Metaplasia, Squamous Respiratory Epithelium, Hyperplasia Lung	1 [1.0] 1 [1.0] 3 [1.7] (50)	2 [1.5] (50)	1 [1.0] (50)	2 [2.0] 1 [1.0] (50)	

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Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS MALE	CONTROL	250 PPM	500 PPM	1000 PPM	
Congestion	1 [3.0]				
Hemorrhage	3 [2.0]	7 [2.6]	3 [2.3]		
Inflammation			1 [2.0]		
Inflammation, Suppurative	1 [1.0]	1 [4.0]			
Inflammation, Granulomatous		1 [1.0]			
Inflammation, Chronic	3 [2.0]	3 [1.7]	6 [1.3]	5 [2.4]	
Thrombosis		1 [2.0]			
Alveolar Epithelium, Hyperplasia	13 [2.0]	11 [1.5]	10 [2.1]	10 [2.3]	
Alveolar Epithelium, Metaplasia,				1 [4.0]	
Squamous					
Alveolar Epithelium, Metaplasia, Mucous				1 [3.0]	
Alveolus, Emphysema				1 [2.0]	
Alveolus, Foreign Body		1 [2.0]			
Alveolus, Infiltration Cellular, Histiocyte	9 [1.9]	7 [1.4]	4 [1.8]	14 [1.8]	
Alveolus, Proteinosis	- 1 - 1		,	1 [3.0]	
Artery, Mineralization	2 [2.5]			11	
Artery, Thrombosis	1 [3.0]				
Bronchiole, Hyperplasia	11	1 [2.0]			
Bronchiole, Inflammation, Chronic		1 [2.0]	1 [2.0]		
Interstitium, Fibrosis		2 [2.0]	1 [2.0]	1 [3.0]	
Nose	(50)	(50)	(49)	(50)	
Foreign Body	5	5	`3 ´	4	
Hyperplasia, Basal Cell		-	-	1 [1.0]	
Inflammation, Suppurative	7 [1.4]	8 [2.4]	8 [1.9]	6 [2.2]	
Inflammation, Chronic	5 [1.6]	1 [1.0]	- []	- []	
Epithelium, Nasolacrimal Duct,	- 1 -1	2 [2.0]	1 [2.0]		
Metaplasia, Squamous		,	,		
Glands, Olfactory Epithelium,	1 [1.0]		1 [2.0]	1 [2.0]	
Hyperplasia	1		1	1 -1	
Glands, Respiratory Epithelium,	1 [4.0]	3 [3.0]	2 [3.0]	2 [3.0]	
Dilatation	. [•]	- []	_[]	-[5.0]	
Glands, Respiratory Epithelium,			2 [2.0]	4 [2.0]	
Hyperplasia			_ [=.0]	. [=.0]	
Goblet Cell, Olfactory Epithelium,			1 [2.0]		
Hyperplasia			. [2.0]		
Goblet Cell, Hyperplasia	3 [1.7]	11 [2.3]	7 [2.3]	5 [2.0]	
Nasolacrimal Duct, Inflammation,	2 [2.0]	2 [2.0]	3 [2.3]	0 [2.0]	
Suppurative	ک <sub>ل</sub> ک.∪ <sub>ا</sub>	ر اِح.∪ا	المارك والمارك		
Olfactory Epithelium, Degeneration			1 [1.0]		
			1 [2.0]		

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS MALE	CONTROL	250 PPM	500 PPM	1000 PPM	
Hyaline					
Olfactory Epithelium, Hyperplasia, Basal Cell		19 [1.1]	27 [1.0]	26 [1.0]	
Olfactory Epithelium, Inflammation, Chronic				1 [2.0]	
Olfactory Epithelium, Metaplasia Olfactory Epithelium, Necrosis	7 [1.7]	4 [1.3] 1 [1.0]	5 [2.4]	5 [2.2] 1 [1.0]	
Olfactory Epithelium, Ulcer Respiratory Epithelium, Degeneration,	1 [1.0]	1 [1.0]	2 [4.0]		
Hyaline Respiratory Epithelium, Hyperplasia Respiratory Epithelium, Inflammation,		15 [2.0]	16 [2.9] 1 [2.0]	23 [2.7] 2 [1.5]	
Chronic Respiratory Epithelium, Necrosis Squamous Epithelium, Hyperplasia Squamous Epithelium, Inflammation Vomeronasal Organ, Inflammation, Suppurative	1 [2.0]		1 [1.0] 1 [1.0]	1 [1.0]	
Pleura Inflammation, Chronic	(5) 4 [1.3]	(3) 3 [1.3]	(5) 4 [1.0]	(6) 6 [1.5]	
Mesothelium, Hyperplasia Trachea Epithelium, Hyperplasia	1 [4.0] (50)	(50)	(50)	(50) 1 [2.0]	
Glands, Cyst		1 [3.0]		. [2.0]	
SPECIAL SENSES SYSTEM					
Eye Degeneration Inflammation, Chronic Active	(50)	(50) 1 [4.0] 1 [4.0]	(49)	(50)	
Bilateral, Lens, Cataract Bilateral, Retina, Atrophy Ciliary Body, Iris, Inflammation, Suppurative	1 [2.0]	1 [4.0] 1 [4.0] 1 [2.0]	1 [2.0] 1 [3.0]	1 [1.0]	
Cornea, Inflammation, Suppurative Cornea, Inflammation, Chronic	2 [1.5]			1 [1.0]	
Lens, Cataract Retina, Atrophy	5 [2.4] 4 [2.5]	1 [2.0]	2 [2.0]	4 [2.5] 2 [3.0]	
Harderian Gland	(50)	(50)	(50)	(50)	

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

Lab: BNW

FISCHER 344 RATS MALE	CONTROL	250 PPM	500 PPM	1000 PPM	
Inflammation, Suppurative	4 [4 0]	1 [3.0]			
Inflammation, Chronic Zymbal's Gland	1 [1.0] (1)	1 [3.0] (0)	(0)	(2)	
JRINARY SYSTEM					
Kidney	(50)	(50)	(50)	(50)	
Atrophy				1 [4.0]	
Infarct		1 [4.0]	2 [3.0]	1 [2.0]	
Infarct, Multiple	47 [0 0]	1 [4.0]	47 [0.0]	FO [O 7]	
Nephropathy Bilateral, Renal Tubule, Cyst	47 [2.3]	47 [2.6]	47 [2.9]	50 [2.7]	
Bilateral, Infarct		1 [3.0]	ı		
Capsule, Dilatation		1 [3.0]	1 [3.0]		
Glomerulus, Inflammation, Suppurative		1 [4.0]	1 [3.0]		
Papilla, Mineralization	5 [1.0]	35 [1.7]	44 [2.1]	41 [2.1]	
Pelvis, Transitional Epithelium,	3 [1.7]	5 [1.8]	14 [2.4]	15 [2.0]	
Hyperplasia	0[]	0[0]	[=]	.0 [2.0]	
Pelvis, Dilatation			1 [4.0]	1 [4.0]	
Renal Tubule, Accumulation, Hyaline			1 [4.0]	1 [4.0]	
Droplet					
Renal Tubule, Cyst		1	3	2	
Renal Tubule, Hyperplasia		3 [3.3]	8 [2.6]	6 [2.2]	
Renal Tubule, Hypertrophy				1 [3.0]	
Renal Tubule, Mineralization	1 [4.0]				
Ureter	(0)	(1)	(0)	(1)	
Urethra	(0)	(0)	(1)	(1)	
Transitional Epithelium, Hyperplasia			1 [3.0]		
Urinary Bladder	(50)	(50)	(49)	(50)	
Calculus Gross Observation			1		
Hemorrhage		1 [2.0]	4 [0.0]		
Inflammation, Chronic	4 [0 0]		1 [2.0]	0.[4.5]	
Transitional Epithelium, Hyperplasia	1 [3.0]		4 [2.0]	2 [1.5]	

\*\*\* END OF MALE \*\*\*

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS FEMALE	CONTROL	250 PPM	500 PPM	1000 PPM
Disposition Summary				
Animals Initially in Study	50	50	50	50
Early Deaths				
Moribund Sacrifice	23	18	17	15
Natural Death	6	5	2	3
Survivors				
Terminal Sacrifice	21	27	31	32
Animals Examined Microscopically	50	50	50	50
ALIMENTARY SYSTEM				
Intestine Large, Cecum	(46)	(45)	(50)	(49)
Necrosis	(12)	(10)	()	1 [3.0]
Intestine Large, Colon	(48)	(46)	(50)	(50)
Artery, Inflammation, Chronic	( - /	( - /	()	1 [3.0]
Intestine Large, Rectum	(48)	(46)	(50)	(50)
Liver	(50)	(50)	(50)	(50)
Angiectasis	1 [3.0]	, ,	2 [2.5]	` ,
Basophilic Focus	27	31	36	32
Clear Cell Focus	15	9	6	5
Eosinophilic Focus				1
Hepatodiaphragmatic Nodule	6 [4.0]	6 [4.0]	4 [4.0]	6 [4.0]
Inflammation, Granulomatous	1 [2.0]		_	
Mixed Cell Focus		,	2	1
Necrosis	4.4.50.41	1 [2.0]	1 [3.0]	3 [3.7]
Vacuolization Cytoplasmic	11 [3.1]	4 [3.0]	4 [2.8]	2 [3.0]
Bile Duct, Hyperplasia		4 [4 0]	1 [3.0]	
Centrilobular, Congestion	4 [0 0]	1 [4.0]		
Periportal, Vacuolization Cytoplasmic Mesentery	1 [2.0] (18)	(18)	(17)	(9)
Necrosis	18 [3.1]	16 [3.0]	(17) 17 [2.9]	8 [3.0]
Oral Mucosa	(0)	(0)	(1)	6 [3.0] (1)
Pancreas	(50)	(49)	(50)	(50)
Fibrosis	(00)	1 [4.0]	(00)	(00)
Hemorrhage		1 [4.0]		
Thrombosis		r -1		1 [4.0]
Artery, Inflammation, Chronic				1 [4.0]

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS FEMALE	CONTROL	250 PPM	500 PPM	1000 PPM	
Stomach, Forestomach	(50)	(50)	(50)	(50)	
Erosion	(00)	(00)	1 [3.0]	(65)	
Inflammation, Suppurative			[1	1 [4.0]	
Inflammation, Chronic		1 [2.0]	1 [2.0]		
Ulcer	5 [3.0]	4 [4.0]	3 [2.3]	1 [4.0]	
Epithelium, Cyst			1 [4.0]		
Epithelium, Erosion		1 [3.0]			
Epithelium, Hyperplasia	11 [1.7]	13 [2.2]	12 [1.9]	8 [1.4]	
Submucosa, Fibrosis	(50)	(40)	1 [3.0]	(50)	
Stomach, Glandular	(50)	(49)	(50)	(50)	
Erosion Ulcer	1 [2.0]		1 [4.0]		
Epithelium, Hyperplasia			1 [4.0]	1 [3.0]	
Tongue	(1)	(2)	(2)	(3)	
Epithelium, Hyperplasia	1 [3.0]	2 [3.0]	2 [2.5]	3 [3.0]	
Tooth	(0)	(2)	(0)	(0)	
Peridontal Tissue, Inflammation	( )	2 [3.5]	( )	,	
CARDIOVASCULAR SYSTEM  Heart	(50)	(50)	(50)	(50)	
Cardiomyopathy Inflammation, Chronic	1 [2.0]	2 [1.5]	2 [2.0]	4 [2 0]	
Atrium, Thrombosis		2 [3.0]	2 [4.0]	1 [2.0] 1 [4.0]	
ENDOCRINE SYSTEM					
Adrenal Cortex Hemorrhage	(50) 1 [3.0]	(49)	(50)	(50)	
Hyperplasia	9 [2.8]	9 [2.8]	13 [2.8]	12 [2.0]	
Necrosis	9 [2.6] 2 [2.5]	J [Z.U]	10 [2.0]	12 [2.0] 1 [3.0]	
Vacuolization Cytoplasmic	11 [2.1]	18 [2.6]	16 [2.4]	7 [2.7]	
Adrenal Medulla	(50)	(49)	(50)	(50)	
Islets, Pancreatic	(50)	(49)	(50)	(50)	
Parathyroid Gland	(49)	(45)	(48)	(47)	
Hyperplasia		1 [3.0]			
Pituitary Gland	(50)	(50)	(50)	(49)	
Cyst	7 [3.3]	2 [3.5]	2 [2.0]	2 [4.0]	
Hemorrhage	2 [4.0]	2 [4.0]	2 [3.5]		

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS FEMALE	CONTROL	250 PPM	500 PPM	1000 PPM	
Hyperplasia Thyroid Gland	10 [3.0] (50)	13 [3.2] (49)	6 [2.7] (50)	16 [2.8] (50)	
Últimobranchial Cyst	1 [1.0]	(1-)	(5-5)	()	
C-cell, Hyperplasia Follicular Cell, Hyperplasia	13 [1.5]	13 [1.7]	15 [1.8] 1 [4.0]	15 [2.2]	
GENERAL BODY SYSTEM					
Peritoneum	(0)	(0)	(1)	(0)	
GENITAL SYSTEM					
Clitoral Gland Cyst	(50)	(49) 2 [3.0]	(50) 2 [3.5]	(50) 1 [1.0]	
Hyperplasia Inflammation, Chronic	2 [3.5]	5 [3.6] 1 [4.0]	7 [3.3] 1 [4.0]	5 [3.8] 3 [3.7]	
Ovary	(50)	(49)	(50)	(50)	
Cyst Cyst, Multiple	3 [3.3]	7 [3.9]	7 [3.3] 1 [4.0]	6 [2.8]	
Uterus	(50)	(49)	(50)	(50)	
Hemorrhage	1 [4.0]	1 [4.0]	4 [3.8]	1 [4.0]	
Pigmentation	4 [4 5]	7 [0 4]	1 [3.0]	E [0 0]	
Endometrium, Hyperplasia Vagina	4 [1.5] (2)	7 [2.4] (0)	2 [2.0] (0)	5 [2.2] (0)	
Epithelium, Hyperplasia, Adenomatous	1 [4.0]	(6)	(0)	(0)	
HEMATOPOIETIC SYSTEM					
Lymph Node Pancreatic, Infiltration Cellular, Histiocyte	(3)	(3)	(5) 1 [4.0]	(2)	
Pancreatic, Pigmentation			1 [3.0]		
Lymph Node, Bronchial	(7)	(9)	(8)	(5)	
Ectasia Hemorrhage	2 [3.0]			1 [4.0]	
Hyperplasia, Lymphoid		1 [3.0]		1 [4.0] 1 [3.0]	
Inflammation	1 [2.0]				
Lymph Node, Mandibular Ectasia	(4)	(1)	(1)	(4) 1 [4.0]	

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Regsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS FEMALE	CONTROL	250 PPM	500 PPM	1000 PPM	
Hyperplasia, Lymphoid	(20)	(00)	(00)	1 [4.0]	
Lymph Node, Mediastinal Angiectasis	(30)	(28)	(26) 1 [4.0]	(31)	
Hemorrhage			1 [3.0]	1 [4.0]	
Hyperplasia, Histiocytic			1 [4.0]		
Hyperplasia, Lymphoid			1 [4.0]	2 [3.5]	
Inflammation, Suppurative Lymph Node, Mesenteric	(40)	(40)	(40)	1 [2.0] (50)	
Ectasia	(48)	(49)	(49) 1 [4.0]	(50)	
Hemorrhage	2 [3.0]		1 [4.0]		
Hyperplasia, Lymphoid	11	1 [2.0]			
Infiltration Cellular, Histiocyte		1 [4.0]			
Necrosis	(50)	(50)	(50)	1 [3.0]	
Spleen Atrophy	(50)	(50)	(50)	(50) 1 [4.0]	
Fibrosis			3 [3.3]	1 [4.0]	
Hematopoietic Cell Proliferation		1 [4.0]	2 [4.0]	2 [4.0]	
Necrosis				2 [4.0]	
Capsule, Fibrosis Thymus	1 [2.0]	(46)	(50)	(50)	
Hyperplasia, Tubular	(48)	(46)	(50) 1 [4.0]	(50)	
NTEGUMENTARY SYSTEM					
Mammary Gland	(50)	(50)	(50)	(50)	
Galactocele	3 [3.0]	()	()	2 [2.5]	
Hyperplasia	1 [2.0]				
Inflammation, Suppurative		1 [4.0]	4 [4 0]	4 [4 0]	
Epithelium, Hyperplasia Skin	(50)	(50)	1 [4.0] (50)	1 [4.0] (50)	
Cyst Epithelial Inclusion	(00)	1 [4.0]	(00)	(00)	
Hyperkeratosis	1 [2.0]	1 [3.0]		1 [2.0]	
Inflammation, Chronic		0.14.03		1 [3.0]	
Ulcer Sebaceous Gland, Hemorrhage		3 [4.0]		2 [3.5]	
Subcutaneous Tissue, Fibrosis		1 [3.0]		1 [2.0]	
Subcutaneous Tissue, Inflammation,		1 [4.0]		. [2.0]	
Suppurative		r -1			

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS FEMALE	CONTROL	250 PPM	500 PPM	1000 PPM	
MUSCULOSKELETAL SYSTEM					
Bone	(50)	(50)	(50)	(50)	
Hyperostosis Maxilla, Fracture		1 [4.0]	1 [4.0]		
Maxilla, Inflammation, Chronic Active			. [•]	1 [3.0]	
Skeletal Muscle	(1)	(0)	(1)	(1)	
IERVOUS SYSTEM					
Brain	(50)	(49)	(50)	(50)	
Compression Demyelination	12 [3.7]	9 [3.6]	4 [4.0]	5 [3.6]	
Gliosis				1 [3.0] 1 [3.0]	
Hemorrhage	1 [4.0]	3 [3.3]	2 [3.5]	3 [3.0]	
Inflammation, Chronic Active Necrosis	1 [4 0]			1 [3.0]	
Thrombosis	1 [4.0]	1 [2.0]	1 [4.0]		
Medulla, Meninges, Inflammation,	1 [2.0]	. []	. []		
Suppurative Medulla, Neuron, Necrosis				4 [2 0]	
Neuron, Degeneration				1 [3.0] 1 [3.0]	
Spinal Cord	(0)	(1)	(1)	(1)	
Gliosis	, ,	. ,	. ,	1 [3.0]	
Inflammation, Chronic Active				1 [3.0]	
RESPIRATORY SYSTEM					
Larynx	(50)	(49)	(50)	(50)	
Foreign Body	2 [4.0]	1 [4.0]	2 [4.0]		
Inflammation, Suppurative	3 [2.0]	3 [2.0]	4 [1.5]	1 [1.0]	
Inflammation, Chronic	2 [1.5]	4 [4 0]	2 [1.5]	2 [1.5]	
Respiratory Epithelium, Hyperplasia Lung	(50)	1 [1.0] (50)	(50)	1 [1.0] (50)	
Cyst, Squamous	(30)	(30)	(30)	(30)	
Hemorrhage	2 [2.0]	1 [1.0]	2 [2.0]	·	
Inflammation, Suppurative	1 [1.0]				
Inflammation, Chronic	11 [1.6]	10 [1.5]	8 [1.8]	16 [1.9]	
Thrombosis			1 [4.0]		

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)

Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

#### P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

FISCHER 344 RATS FEMALE	CONTROL	250 PPM	500 PPM	1000 PPM
Alveolar Epithelium, Hyperplasia Alveolar Epithelium, Metaplasia, Squamous	3 [2.7]	1 [1.0]	5 [2.4]	7 [2.1] 1 [4.0]
Alveolus, Infiltration Cellular, Histiocyte Alveolus, Proteinosis	16 [1.1]	6 [1.3]	11 [1.6]	14 [1.6] 2 [2.5]
Bronchiole, Hyperplasia Bronchiole, Inflammation, Chronic Interstitium, Fibrosis	1 [4.0] 1 [2.0]	3 [2.3] 1 [3.0]		1 [2.0] 3 [2.0]
Nose Foreign Body Hyperplasia, Basal Cell	(50)	(48) 4	(50) 2	(50) 3 1 [1.0]
Inflammation, Suppurative Inflammation, Chronic Inflammation, Chronic Active	1 [3.0] 1 [1.0]	3 [3.3] 3 [2.0]	1 [2.0] 4 [1.3] 1 [4.0]	2 [1.5] 1 [2.0]
Glands, Respiratory Epithelium, Dilatation Goblet Cell, Hyperplasia	1 [3.0] 4 [1.0]	6 [2.3]	1 [4.0]	5 [2.0]
Nasolacrimal Duct, Inflammation, Suppurative	4 [2.3]	3 [2.0]	2 [2.5]	1 [3.0]
Nasolacrimal Duct, Inflammation, Chronic Olfactory Epithelium, Degeneration, Hyaline	1 [3.0] 2 [1.5]	1 [2.0]		
Olfactory Epithelium, Hyperplasia Olfactory Epithelium, Hyperplasia, Basal Cell		14 [1.0]	1 [1.0] 25 [1.0]	31 [1.1]
Olfactory Epithelium, Inflammation, Granulomatous		1 [3.0]		
Olfactory Epithelium, Metaplasia Olfactory Epithelium, Necrosis Respiratory Epithelium, Degeneration,	1 [1.0]	1 [3.0]	1 [1.0] 3 [1.0]	2 [2.0]
Hyaline Respiratory Epithelium, Hyperplasia	. [9]		4 [3.0]	6 [2.3]
Respiratory Epithelium, Metaplasia, Squamous Turbinate, Necrosis		1 [1.0]	1 [4.0]	
Pleura Inflammation, Chronic	(15) 15 [1.4]	(16) 15 [1.2]	(16) 14 [1.4]	(20) 20 [1.8]
Mesothelium, Hyperplasia Trachea Glands, Cyst	(50)	1 [1.0] (49)	(50) 1 [2.0]	(50)

a - Number of animals examined microscopically at site and number of animals with lesion

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Species/Strain: RATS/F 344

Route: RESPIRATORY EXPOSURE WHOLE BODY

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

CUMENE

**CAS Number:** 98-82-8

Pathologist: PIERCE, J. - RENNE, R.

Date Report Reqsted: 01/12/2006

Time Report Reqsted: 09:48:09 First Dose M/F: 06/04/01 / 06/04/01

Lab: BNW

FISCHER 344 RATS FEMALE	CONTROL	250 PPM	500 PPM	1000 PPM	
SPECIAL SENSES SYSTEM					
Eye	(50)	(49)	(50)	(50)	
Bilateral, Lens, Cataract	1 [4.0]				
Bilateral, Retina, Atrophy	1 [3.0]	1 [2.0]	1 [3.0]	4 [0.0]	
Cornea, Epithelium, Hyperplasia				1 [2.0]	
Cornea, Inflammation Cornea, Inflammation, Suppurative	1 [4.0]			1 [1.0]	
Lens, Cataract	5 [2.2]	2 [3.0]	5 [3.0]	3 [2.0]	
Retina, Atrophy	3 [2.0]	5 [2.2]	8 [2.1]	2 [2.0]	
Harderian Gland	(50)	(49)	(50)	(50)	
Inflammation, Chronic	1 [1.0]	( - /	1 [2.0]	1 [1.0]	
Zymbal's Gland	(1)	(0)	(1)	(1)	
JRINARY SYSTEM					
Kidney	(50)	(50)	(50)	(50)	
Infarct, Multiple		, ,	1 [2.0]		
Nephropathy	38 [1.4]	37 [1.5]	41 [1.9]	44 [1.9]	
Artery, Inflammation, Chronic Active	0.14.03	0.14.03	4.54.03	1 [2.0]	
Papilla, Mineralization	6 [1.0]	3 [1.3]	4 [1.0]	6 [1.0]	
Pelvis, Transitional Epithelium, Hyperplasia	1 [1.0]	1 [2.0]	6 [1.3]	1 [1.0]	
Pelvis, Transitional Epithelium,	23 [1.3]	27 [1.1]	27 [1.1]	22 [1.3]	
Mineralization	20 [1.0]	21 [1.1]	21 [1.1]	22 [1.0]	
Pelvis, Dilatation	1 [2.0]				
Renal Tubule, Accumulation, Hyaline	1 [4.0]		1 [4.0]		
Droplet					
Renal Tubule, Cyst	1		1	1	
Renal Tubule, Pigmentation	1 [4.0]	(40)	(50)	(50)	
Urinary Bladder Transitional Epithelium, Hyperplasia	(50)	(49)	(50)	(50)	
rransilionai ⊏pilitellum, ⊓yperpiasia	1 [2.0]	2 [2.0]	2 [2.0]	1 [2.0]	

\*\*\* END OF REPORT \*\*\*

a - Number of animals examined microscopically at site and number of animals with lesion

b - Average severity grade (1-minimal; 2-mild; 3-moderate; 4-marked)